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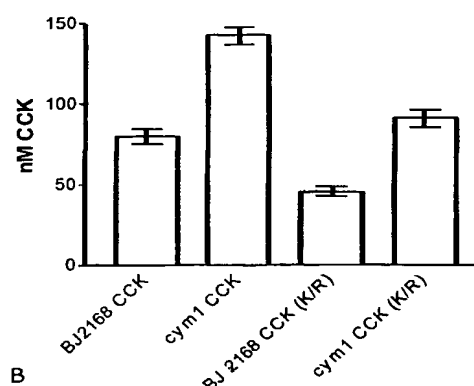
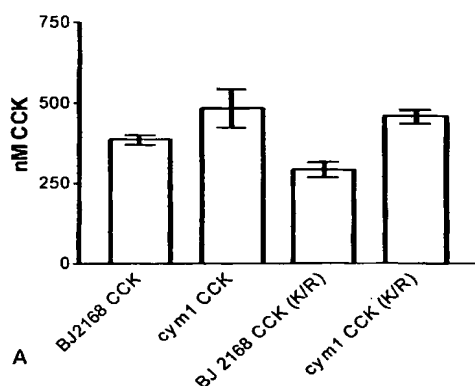
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(54) Title: METHODS FOR INCREASING THE PRODUCTION OF A RECOMBINANT POLYPEPTIDE FROM A HOST CELL



(57) Abstract: The present invention provides a host cell comprising a nucleic acid sequence encoding a recombinant polypeptide in which the production of a naturally occurring metalloprotease comprising the sequence HXXEH has been reduced or inhibited by genetic manipulation. The present invention also relates to methods for enhancing the production of a polypeptide from a cell by disrupting the synthesis or activity of the metalloprotease. In particular, the present invention relates to methods for enhancing the secretion of recombinant polypeptides from host cells such as, but not limited to, yeast and bacterial cells. The metalloprotease is a member of the pepsin family of proteases, characterized by comprising the sequence HXXEH, where X is any amino acid.

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INTERNATIONAL SEARCH REPORT

International Application No
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A. CLASSIFICATION OF SUBJECT MATTER
IPC 7 C12N15/57 C12P21/02 C12N1/19

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 C12P C12N

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, BIOSIS, WPI Data, PAJ, SEQUENCE SEARCH

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 99 33960 A (ESTELL DAVID A ; GENENCOR INT (US)) 8 July 1999 (1999-07-08) claims 1-8 page 2, line 18 - line 26 ---	1-22
X	WO 97 46689 A (GIST BROCADES BV ; DEN HOMBERGH JOHANNES PETRUS T (NL); VISSER JACO) 11 December 1997 (1997-12-11) cited in the application claims page 15, paragraph 5.4 ---	1-22
X	WO 96 29391 A (NOVONORDISK AS ; LEHMBECK JAN (DK)) 26 September 1996 (1996-09-26) claims ---	1-22

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☒ Further documents are listed in the continuation of box C.

☒ Patent family members are listed in annex.

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Date of the actual completion of the international search

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INTERNATIONAL SEARCH REPORT

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C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	<p>EGEL-MITANI M ET AL: "Yield improvement of heterologous peptides expressed in yps1-disrupted <i>Saccharomyces cerevisiae</i> strains"</p> <p>ENZYME AND MICROBIAL TECHNOLOGY, vol. 26, no. 9-10, June 2000 (2000-06), pages 671-677, XP002283345</p> <p>ISSN: 0141-0229</p> <p>abstract</p> <p style="text-align: center;">-----</p>	1-22

INTERNATIONAL SEARCH REPORT

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